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De	A PRINCIPLE OF HER 2004 OF A READ THORITY						
	To: DAVID P. HALSTEAD ROPES & GRAY LLP	PCT					
Po	ONE INTERNATIONAL PLACE BOSTON, MA 02110-2624 DES & Gray	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT					
	nbol #: <u>Cwrv-Pwo.043</u>	OR THE DECLARATION					
	tion Due: Aile Article 19 Amendment	(PCT Rule 44.1)					
De	adline(s): 04 Feb 2004/09 How 2004	Date of Mailing (day/month/year)					
	Applicant's or agent's file reference CWRU-PWO-043	FOR FURTHER ACTION See paragraphs 1 and 4 below					
· · .	International application No. PCT/US03/23082	International filing date (day/month/year) 24 July 2003 (24 07 2003)					
	Applicant_ CASE WI_TERN RESERVE UNIVERSITY						
	The applicant is hereby notified that the international search has been established and is trans: herewise. Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the international application (see Rule): When? The time limit for filing such amendments is normally two months from the date of transmittal of the international search report. Where? Directly to the International Bureau of WIPO, 34, chemin des Colombettes . 1211 Geneva 20, Switzerland, Facsimile No.: (41-22) 740.14.35 For more detailed instructions, see the notes on the accompanying sheet. 2. The applicant is hereby notified that no international search report will be established and that the declaration under Article 17(2)(a) to that effect is transmitted herewith. 3. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that: the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices. no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made. 4. Reminders Shortly after 18 months from the priority date, the international application will be published by the International Bureau applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority c must reach the International Bureau as provided in Rules 90 bis.1 and 90 bis.3, respectively, before the completion of the tecl preparations for international publication. Within 19 months from the priority date, but only in respect of some designated Offices, a demand for international prelim examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date, in some Offices even later), otherwise the applicant must, within 20 months						
· [Guide, Volume II, National Chapters and the WIPO Internet sit Name and mailing address of the ISA/US	Authorized officer					
	Mail Stop PCT, Attn: ISA/US Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450	Cherre Bell-Harris An					

Facsimile No. (703)305-3230
Form PCT/ISA/220 (April 2002)

(See notes on accompanying sheet)

PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY To: DAVID P. HALSTEAD **ROPES & GRAY LLP** ONE INTERNATIONAL PLACE NOTIFICATION OF TRANSMITTAL OF BOSTON, MA 02110-2624 THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION (PCT Rule 44.1) Date of Mailing 09 JAN 2004 (day/month/year) Applicant's or agent's file reference CWRU-PWO-043 FOR FURTHER ACTION See paragraphs 1 and 4 below International application No. International filing date PCT/US03/23082 (day/month/year) Applicant CASE WESTERN RESERVE UNIVERSITY The applicant is hereby notified that the international search report has been established and is transmitted herewith. Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the international application (see Rule 46): The time limit for filing such amendments is normally two months from the date of transmittal of the international search report. Where? Directly to the International Bureau of WIPO, 34, chemin des Colombettes 1211 Geneva 20, Switzerland, Facsimile No.: (41-22) 740.14.35 For more detailed instructions, see the notes on the accompanying sheet. The applicant is hereby notified that no international search report will be established and that the declaration under Article 17(2)(a) to that effect is transmitted herewith. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that: 3 the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices. no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made. Reminders Shortly after 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in Rules 90 bis.1 and 90 bis.3, respectively, before the completion of the technical preparations for international publication. Within 19 months from the priority date, but only in respect of some designated Offices, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later); otherwise the applicant must, within 20 months from the priority date, perform the prescribed acts for entry into the national phase before those designated Offices. In respect of other designated Offices, the time limit of 30 months (or later) will apply even if no demand is filed within 19 months. See the Annex to Form PCT/IB/301 and, for details about the applicable time limits, Office by Office, see the PCT Applicant's Guide, Volume II, National Chapters and the WIPO Internet site. Name and mailing address of the ISA/US Authorized officer Mail Stop PCT, Atm: ISA/US Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450 Telephone No. (703) 308-1235 Facsimile No. (703)305-3230

(See notes on accompanying sheet)

Form PCT/ISA/220 (April 2002)

PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH-REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference CWRU-PWO-043	FOR FURTHER ACTION		e Notification of Transmittal of International Search eport (Form PCT/ISA/220) as well as, where applicable, em 5 below.				
International application No. PCT/US03/23082	International filing date (day/mod 24 July 2003 (24.07.2003)	nth/year)	(Earliest) Priority Date (day/month/year)				
Applicant CASE WESTERN RESERVE UNIVERSITY							
This international search report has been prepared by this International Searching Authority and is transmitted to the applicant according to Article 18. A copy is being transmitted to the International Bureau.							
This international search report consists of a total of _3_ sheets. It is also accompanied by a copy of each prior art document cited in this report.							
Basis of the Report a. With regard to the language, the international search was carried out on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.							
the international search was carried out on the basis of a translation of the international application furnished to this Authority (Rule 23.1(b)). b. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international search was carried out on the basis of the sequence listing:							
contained in the interna	tional application in written form.						
filed together with the i	filed together with the international application in computer readable form.						
furnished subsequently	to this Authority in written form.		•• .				
furnished subsequently	to this Authority in computer readable	form.					
the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.							
the statement that the ir been furnished.	nformation recorded in computer reada	ble form is ic	dentical to the written sequence listing has				
2. Certain claims were for	ound unsearchable (See Box I).	*					
3. Unity of invention is la 4. With regard to the title,	acking (See Box II).						
the text is approved as	submitted by the applicant.						
the text has been estable Please See Continuation Sheet	ished by this Authority to read as follo	ws:					
5. With regard to the abstract,		•	•				
the text is approved as	submitted by the applicant.	•					
			as it appears in Box III. The applicant may, ort, submit comments to this Authority.				
6. The figure of the drawings to be	6. The figure of the drawings to be published with the abstract is Figure No						
as suggested by the app	· · · · · · · · · · · · · · · · · · ·		None of the figures				
	illed to suggest a figure.						
	er characterizes the invention.						
	•						

Form PCT/ISA/210 (first sheet) (July 1998)

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/23082

A. CLASSIFICATION OF SUBJECT MATTER IPC(7) : A61K 31/425, 31/50, 31/497, 31/44, 31/42 US CL : 514/369, 252.05, 254.02, 256, 342, 375, 376							
According to	According to International Patent Classification (IPC) or to both national classification and IPC						
B. FIELI	DS SEARCHED	•					
Minimum documentation searched (classification system followed by classification symbols) U.S.: 514/369, 252.05, 254.02, 256, 342, 375, 376							
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched							
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Please See Continuation Sheet							
C. DOCU	JMENTS CONSIDERED TO BE RELEVANT						
Category *	Citation of document, with indication, where ap	opropriate, of the relevant passages	Relevant to claim No.				
X	HIRSCH et al. The Role of Glial Reaction and Inflar		1-9				
	2003, Volume 991, pages 214-228, especially page 2 WO 02/49626 A2 (SMITHKLINE BEEECHAM P.I		1-8				
X 	entire document, especially page 2, lines 1-40, page						
A			9				
х	BRIEDERT et al. Protective Action of the Peroxisome Proliferator-Activated Receptor-		1-9				
	Gamma Agonist Pioglitazone in a Mouse Model of Parkinson's Disease, 2002, Journal of Neurochemistry, Volume 82, Pages 615-624. See entire document, especially page 615, column 1 to page 616, column 1 and page 621, column 1 to column 2.+.						
x	US 6,369,098 B1 (PERSHADSINGH et al) 09 April		1-8				
Y .	column 12, lines 24-37, column 20, line 23 to column	in 24, line 34, column 27, mies 26-37.	9				
	· · · · · · · · · · · · · · · · · · ·						
Further	documents are listed in the continuation of Box C.	See patent family annex.	•				
* Sp	ecial categories of cited documents:	"T" later document published after the inter date and not in conflict with the applic					
	defining the general state of the art which is not considered to be ar relevance	principle or theory underlying the inve					
"E" earlier application or patent published on or after the international filing date		"X" document of particular relevance; the considered novel or cannot be consider when the document is taken alone					
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)		"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is					
"O" document	referring to an oral disclosure, use, exhibition or other means	combined with one or more other such being obvious to a person skilled in the					
	published prior to the international filing date but later than the tte claimed	"&" document member of the same patent family					
Date of the ac	tual completion of the international search	Date of mailing of the international search	th report				
	2003 (14.12.2003)		ロカ・コトに 人口の				
	iling address of the ISA/US	Authorized officer Authorized Officer Authorized Officer Authorized Officer Authorized Officer					
Com	Stop PCT, Attn: ISA/US unissioner for Patents	Donna Jagoe el el Francisco					
. Alex	. Box 1450 tandria, Virginia 22313-1450 . (703)305-3230	Telephone No. (703) 308-1235	`				

1	PC1/USU3/23U82
INTERNATIONAL SEARCH REPORT	
*.	
Continuation of Item 4 of the first sheet: The title does not meet PCT Rule 4.3 because it is not properly describe the invention.	ntion. The new title is "Methods for the Treatment of
Parkinson's Disease".	
	· .
Continuation of B. FIELDS SEARCHED Item 3: WEST 2.1, STN, file cluster MEDICINE, Search terms, PPAR gamma agonist,	this rolidine dione parkinson's microglial tyrosine
WEST 2.1, STN, file cluster MEDICINE, Search terms, FFAR gamma agomst,	mazonumedione, parameter s, androgram, cyrosmo-
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Form PCT/ISA/210 (second sheet) (July 1998)

NOTESTO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under Article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article," "Rule" and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions, respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international publication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Preliminary Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When? Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been/is filed, see below.

How? Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

"What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- 1. [Where originally there were 48 claims and after amendment of some claims there are 51]: "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers; claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- 2. [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
 "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or
 "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4. [Where various kinds of amendments are made]: "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under Article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international application is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments and any accompanying statement, under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the time of filing the amendments (and any statement) with the International Bureau, also file with the International Preliminary Examining Authority a copy of such amendments (and of any statement) and, where required, a translation of such amendments for the procedure before that Authority (see Rules 55.3(a) and 62.2, first sentence). For further information, see the Notes to the demand form (PCT/IPEA/401).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see the PCT Applicant's Guide, Volume II.